Al-Mustaqbal University College Department of Pharmacy 5th stage/2024 Clinical Toxicology Lecture: 2



Anticholinergic Toxicity

Autonomic nervous system (ANS)

Sympathetic

Dominated by (Adrenaline) (Adrenergic) fight or flight



Parasympathetic

Dominated by Acetylcholine (Cholinergic) Rest and digest



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Outline

Anticholinergic toxicity?

Multiple organ adverse effect

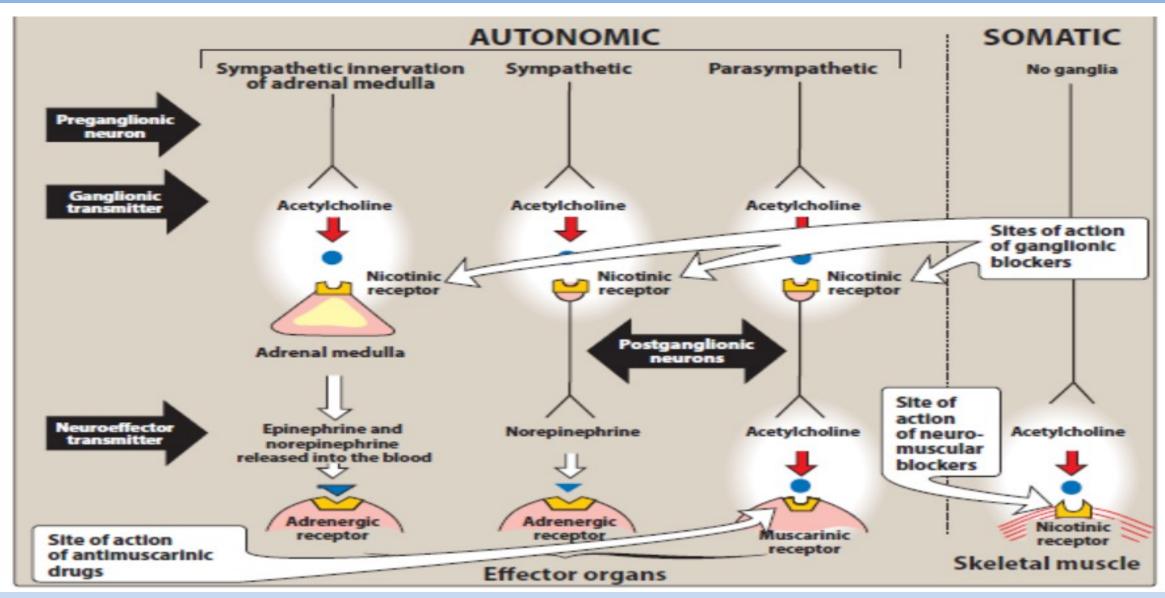
Autonomic Nervous System

Toxidrome Clinical Manifestation:

Clinical Manifestation:

Management and Antidote

Autonomic nervous system



Clinical Toxicology 5th stage

Al-Mustaqbal University

College of Pharmacy

Weaam J. Abbas

Anticholinergic Drug Toxicity

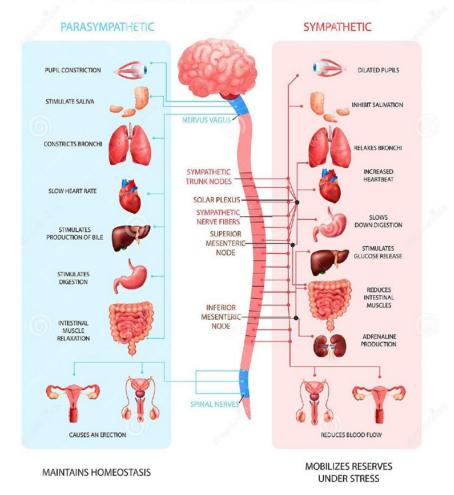
Cholinergic antagonist

is an agents that bind to cholinorceptors (muscarinic or nicotinic) and prevent the effects of acetylcholine (ACh) and other cholinergic agonists.

The most clinically useful of these agents are selective blockers of muscarinic receptors.

The effects of parasympathetic innervation are, thus, interrupted, and the actions of sympathetic stimulation are left unopposed.

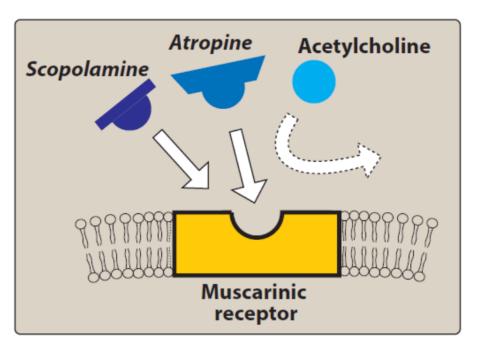
PERIPHERAL AUTONOMIC NERVOUS SYSTEM



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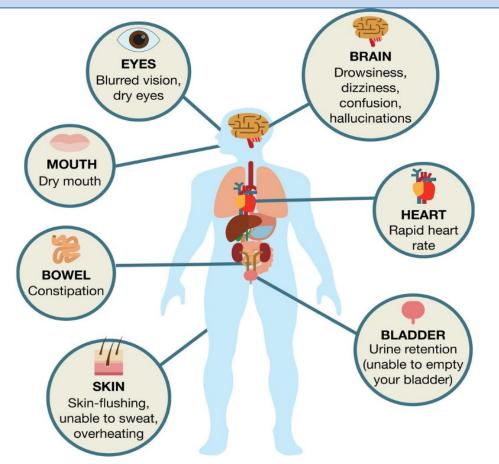
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- Anticholinergic syndrome is produced by the inhibition of cholinergic neurotransmission at muscarinic receptor sites.
- It commonly follows the ingestion of a wide variety of prescription and over-the-counter medications.



Signs and symptoms:

Clinical manifestations are caused by:



CNS effects

Peripheral nervous system effects

Or both

Clinical manifestations may include:

- > Flushing
- > Dry skin and mucous membranes
- > Mydriasis with loss of accommodation
- > Altered mental status
- > Fever

Anticholinergic Toxidrome × Mad as a hatter Altered mental status Blind as a bat Pupillary dilation with loss of accommodation Red as a beet Vasodilation with skin hyperemia Hot as a hare Anhydrosis with temperature elevation Dry as a bone Drying of mucosal surfaces and skin Full as a flask Urinary retention

Stuffed as a pepper

Constipation

Additional manifestations include the following:

- > Sinus tachycardia
- > Decreased bowel sounds
- > Urinary retention
- > Hypertension

Anticholinergic Drugs

Agents with anticholinergic properties are as follows:

- 1. Anticholinergics
- **✓** Atropine
- **✓** scopolamine
- 2. Antihistamines
- **✓** Chlorpheniramine
- **✓** Cyproheptadine
- **✓** Diphenhydramine
- **✓** Promethazine

Anticholinergic Drugs

Agents with anticholinergic properties are as follows:

- 3. Antipsychotics
- **✓** Chlorpromazine
- **✓** Clozapine
- **✓** Thioridazine
- 4. Antispasmodics
- ✓ Clidinium
- **✓** Hyoscyamine
- **✓** Propantheline

Management

Treatment:

- ✓ Initial assessment and stabilization are required
- ✓ Ensure an adequate airway and check that breathing is present and maintained.
- ✓ Assess circulation and initiate cardiac and pulse monitoring.
- ✓ GI decontamination with activated charcoal is recommended.
- ✓ Ipecac syrup is contraindicated.
- ✓ Ventricular arrhythmias can be treated with lidocaine.
- ✓ Manage seizures with benzodiazepines.

Management - Antidote

The antidote for anticholinergic toxicity is

Physostigmine



(Physostigmine salicylate.)

- * Physostigmine is the only reversible acetylcholinesterase inhibitor capable of directly antagonizing the CNS manifestations of anticholinergic toxicity.
- ***It is an uncharged tertiary amine that efficiently crosses the blood brain barrier.**

Management - Antidote

By inhibiting acetylcholinesterase, the enzyme responsible for the hydrolysis of acetylcholine, an increased concentration of acetylcholine augments stimulation at muscarinic and nicotinic receptors.

Physostigmine can reverse the central effects of coma, seizures, severe dyskinesias, hallucinations, agitation, and respiratory depression.

The most common indication for physostigmine is to control <u>agitated delirium</u>.

Physostigmine is contraindicated in patients with cardiac conduction disturbances (prolonged PR and QRS intervals) on ECG analysis.

